

In A.) the the wild-type and the mutated EGFR-II are shown, in B.) the wild type and the mutated MuSK is shown. Both EGFR-II and the mutated MuSK can be used as selectable cell surface markers.

| | MRPSGTAGAA | LLALLAALCP | ASRALEEKKV | CQGTSNKLTQ | LGTFEDHFLS | 50 |
|----|------------|------------|--------------------|------------|------------|------|
| | LQRMFNNCEV | VLGNLEITYV | QRNYDLSFLK | TIQEVAGYVL | IALNTVERIP | 100 |
| | LENLQIIRGN | MYYENSYALA | VLSNYDANKT | GLKELPMRNL | QEILHGAVRF | 150 |
| | SNNPALCNVE | SIQWRDIVSS | DFLSNMSMDF | QNHLGSCQKC | DPSCPNGSCW | 200 |
| 5 | GAGEENCQKL | TKIICAQQCS | GRCRGKSPSD | CCHNQCAAGC | TGPRESDCLV | 250 |
| | CRKFRDEATC | KDTCPPLMLY | NPTTYQMDVN | PEGKYSFGAT | CVKKCPRNYV | 300 |
| | VTDHGSCVRA | CGADSYEMEE | DGVRKCKKCE | GPCRKVCNGI | GIGEFKDSLS | 350 |
| | INATNIKHFK | NCTSISGDLH | ILPVAFRGDS | FTHTPPLDPQ | ELDILKTVKE | 400 |
| | ITGFLLIQAW | PENRTDLHAF | ENLEIIRGRT | KQHGQFSLAV | VSLNITSLGL | 450 |
| 10 | RSLKEISDGD | VIISGNKNLC | YANTINWKKL | FGTSGQKTKI | ISNRGENSCK | 500 |
| | ATGQVCHALC | SPEGCWGPEP | RDCVSCRNVS | RGRECVDKCN | LLEGEPREFV | 550 |
| | ENSECIQCHP | ECLPQAMNIT | CTGRGPDNCI | QCAHYIDGPH | CVKTCPAGVM | 600 |
| | GENNTLVWKY | ADAGHVCHLC | HPNCTYGCTG | PGLEGCPTNG | PKIPSIATGM | 650 |
| | VGALLLLLVV | ALGIGLFMRR | RHIVRKRTLR | RLLQERELVE | PLTPSGEAPN | 700 |
| 15 | QALLRILKET | EFKKIKVLGS | ${\tt GAFGTVYKGL}$ | WIPEGEKVKI | PVAIKELREA | 750 |
| | TSPKANKEIL | DEAYVMASVD | NPHVCRLLGI | CLTSTVQLIT | QLMPFGCLLD | 800 |
| | YVREHKDNIG | SQYLLNWCVQ | IAKGMNYLED | RRLVHRDLAA | RNVLVKTPQH | 850 |
| | VKITDFGLAK | LLGAEEKEYH | AEGGKVPIKW | MALESILHRI | YTHQSDVWSY | 900 |
| | GVTVWELMTF | GSKPYDGIPA | SEISSILEKG | ERLPQPPICT | IDVYMIMVKC | 950 |
| 20 | WMIDADSRPK | FRELIIEFSK | MARDPQRYLV | IQGDERMHLP | SPTDSNFYRA | 1000 |
| | LMDEEDMDDV | VDADEYLIPQ | QGFFSSPSTS | RTPLLSSLSA | TSNNSTVACI | 1050 |
| | DRNGLQSCPI | KEDSFLQRYS | SDPTGALTED | SIDDTFLPVP | EYINQSVPKR | 1100 |
| | PAGSVQNPVY | HNQPLNPAPS | RDPHYQDPHS | TAVGNPEYLN | TVQPTCVNST | 1150 |
| | FDSPAHWAQK | GSHQISLDNP | DYQQDFFPKE | AKPNGIFKGS | TAENAEYLRV | 1200 |
| 25 | APQSSEFIGA | | | | | 1210 |

FIGURE 2

| | ATGCGACCCT | CCGGGACGGC | CGGGGCAGCG | CTCCTGGCGC | TGCTGGCTGC | 50 |
|----|--------------------|------------|------------|------------|------------|------|
| | GCTCTGCCCG | GCGAGTCGGG | CTCTGGAGGA | AAAGAAAGTT | TGCCAAGGCA | 100 |
| | CGAGTAACAA | GCTCACGCAG | TTGGGCACTT | TTGAAGATCA | TTTTCTCAGC | 150 |
| 5 | CTCCAGAGGA | TGTTCAATAA | CTGTGAGGTG | GTCCTTGGGA | ATTTGGAAAT | 200 |
| | TACCTATGTG | CAGAGGAATT | ATGATCTTTC | CTTCTTAAAG | ACCATCCAGG | 250 |
| | AGGTGGCTGG | TTATGTCCTC | ATTGCCCTCA | ACACAGTGGA | GCGAATTCCT | 300 |
| | TTGGAAAACC | TGCAGATCAT | CAGAGGAAAT | ATGTACTACG | AAAATTCCTA | 350 |
| | TGCCTTAGCA | GTCTTATCTA | ACTATGATGC | AAATAAAACC | GGACTGAAGG | 400 |
| 10 | AGCTGCCCAT | GAGAAATTTA | CAGGAAATCC | TGCATGGCGC | CGTGCGGTTC | 450 |
| | AGCAACAACC | CTGCCCTGTG | CAACGTGGAG | AGCATCCAGT | GGCGGGACAT | 500 |
| | AGTCAGCAGT | GACTTTCTCA | GCAACATGTC | GATGGACTTC | CAGAACCACC | 550 |
| | TGGGCAGCTG | CCAAAAGTGT | GATCCAAGCT | GTCCCAATGG | GAGCTGCTGG | 600 |
| | GGTGCAGGAG | AGGAGAACTG | CCAGAAACTG | ACCAAAATCA | TCTGTGCCCA | 650 |
| 15 | GCAGTGCTCC | GGGCGCTGCC | GTGGCAAGTC | CCCCAGTGAC | TGCTGCCACA | 700 |
| | ACCAGTGTGC | TGCAGGCTGC | ACAGGCCCCC | GGGAGAGCGA | CTGCCTGGTC | 750 |
| | TGCCGCAAAT | TCCGAGACGA | AGCCACGTGC | AAGGACACCT | GCCCCCACT | 800 |
| | CATGCTCTAC | AACCCCACCA | CGTACCAGAT | GGATGTGAAC | CCCGAGGGCA | 850 |
| | AATACAGCTT | TGGTGCCACC | TGCGTGAAGA | AGTGTCCCCG | TAATTATGTG | 900 |
| 20 | GTGACAGATC | ACGGCTCGTG | CGTCCGAGCC | TGTGGGGCCG | ACAGCTATGA | 950 |
| | GATGGAGGAA | GACGGCGTCC | GCAAGTGTAA | GAAGTGCGAA | GGGCCTTGCC | 1000 |
| | ${\tt GCAAAGTGTG}$ | TAACGGAATA | GGTATTGGTG | AATTTAAAGA | CTCACTCTCC | 1050 |
| | ATAAATGCTA | CGAATATTAA | ACACTTCAAA | AACTGCACCT | CCATCAGTGG | 1100 |
| | CGATCTCCAC | ATCCTGCCGG | TGGCATTTAG | GGGTGACTCC | TTCACACATA | 1150 |
| 25 | CTCCTCCTCT | GGATCCACAG | GAACTGGATA | TTCTGAAAAC | CGTAAAGGAA | 1200 |
| | ATCACAGGGT | TTTTGCTGAT | TCAGGCTTGG | CCTGAAAACA | GGACGGACCT | 1250 |
| | CCATGCCTTT | GAGAACCTAG | AAATCATACG | CGGCAGGACC | AAGCAACATG | 1300 |
| | GTCAGTTTTC | TCTTGCAGTC | GTCAGCCTGA | ACATAACATC | CTTGGGATTA | 1350 |
| | CGCTCCCTCA | AGGAGATAAG | TGATGGAGAT | GTGATAATTT | CAGGAAACAA | 1400 |

FIGURE 3A

| | AAATTTGTGC | TATGCAAATA | CAATAAACTG | GAAAAAACTG | TTTGGGACCT | 1450 |
|----|--------------------|------------|--------------------|--------------------|------------|------|
| | CCGGTCAGAA | AACCAAAATT | ATAAGCAACA | GAGGTGAAAA | CAGCTGCAAG | 1500 |
| | GCCACAGGCC | AGGTCTGCCA | TGCCTTGTGC | TCCCCGAGG | GCTGCTGGGG | 1550 |
| 5 | CCCGGAGCCC | AGGGACTGCG | TCTCTTGCCG | GAATGTCAGC | CGAGGCAGGG | 1600 |
| | AATGCGTGGA | CAAGTGCAAG | CTTCTGGAGG | GTGAGCCAAG | GGAGTTTGTG | 1650 |
| | GAGAACTCTG | AGTGCATACA | GTGCCACCCA | GAGTGCCTGC | CTCAGGCCAT | 1700 |
| | GAACATCACC | TGCACAGGAC | GGGGACCAGA | CAACTGTATC | CAGTGTGCCC | 1750 |
| | ACTACATTGA | CGGCCCCAC | TGCGTCAAGA | CCTGCCCGGC | AGGAGTCATG | 1800 |
| 10 | GGAGAAAACA | ACACCCTGGT | CTGGAAGTAC | GCAGACGCCG | GCCATGTGTG | 1850 |
| | CCACCTGTGC | CATCCAAACT | GCACCTACGG | ATGCACTGGG | CCAGGTCTTG | 1900 |
| | AAGGCTGTCC | AACGAATGGG | CCTAAGATCC | CGTCCATCGC | CACTGGGATG | 1950 |
| | GTGGGGGCCC | TCCTCTTGCT | GCTGGTGGTG | GCCCTGGGGA | TCGGCCTCTT | 2000 |
| | CATGCGAAGG | CGCCACATCG | TTCGGAAGCG | CACGCTGCGG | AGGCTGCTGC | 2050 |
| 15 | AGGAGAGGGA | GCTTGTGGAG | CCTCTTACAC | CCAGTGGAGA | AGCTCCCAAC | 2100 |
| | CAAGCTCTCT | TGAGGATCTT | GAAGGAAACT | GAATTCAAAA | AGATCAAAGT | 2150 |
| | GCTGGGCTCC | GGTGCGTTCG | GCACGGTGTA | TAAGGGACTC | TGGATÇCCAG | 2200 |
| | AAGGTGAGAA | AGTTAAAATT | CCCGTCGCTA | ${\tt TCAAGGAATT}$ | AAGAGAAGCA | 2250 |
| | ACATCTCCGA | AAGCCAACAA | GGAAATCCTC | GATGAAGCCT | ACGTGATGGC | 2300 |
| 20 | CAGCGTGGAC | AACCCCCACG | TGTGCCGCCT | GCTGGGCATC | TGCCTCACCT | 2350 |
| | CCACCGTGCA | ACTCATCACG | CAGCTCATGC | CCTTCGGCTG | CCTCCTGGAC | 2400 |
| | ${\tt TATGTCCGGG}$ | AACACAAAGA | ${\tt CAATATTGGC}$ | TCCCAGTACC | TGCTCAACTG | 2450 |
| | GTGTGTGCAG | ATCGCAAAGG | GCATGAACTA | CTTGGAGGAC | CGTCGCTTGG | 2500 |
| | TGCACCGCGA | CCTGGCAGCC | AGGAACGTAC | TGGTGAAAAC | ACCGCAGCAT | 2550 |
| 25 | GTCAAGATCA | CAGATTTTGG | GCTGGCCAAA | CTGCTGGGTG | CGGAAGAGAA | 2600 |
| | AGAATACCAT | GCAGAAGGAG | GCAAAGTGCC | TATCAAGTGG | ATGGCATTGG | 2650 |
| | AATCAATTTT | ACACAGAATC | TATACCCACC | AGAGTGATGT | CTGGAGCTAC | 2700 |
| | GGGGTGACCG | TTTGGGAGTT | GATGACCTTT | GGATCCAAGC | CATATGACGG | 2750 |
| | AATCCCTGCC | AGCGAGATCT | CCTCCATCCT | GGAGAAAGGA | GAACGCCTCC | 2800 |

| | CTCAGCCACC | CATATGTACC | ATCGATGTCT | ACATGATCAT | GGTCAAGTGC | 2850 |
|----|------------|--------------------|--------------------|------------|------------|------|
| | TGGATGATAG | ACGCAGATAG | TCGCCCAAAG | TTCCGTGAGT | TGATCATCGA | 2900 |
| | ATTCTCCAAA | ATGGCCCGAG | ACCCCCAGCG | CTACCTTGTC | ATTCAGGGGG | 2950 |
| | ATGAAAGAAT | GCATTTGCCA | AGTCCTACAG | ACTCCAACTT | CTACCGTGCC | 3000 |
| 5 | CTGATGGATG | AAGAAGACAT | GGACGACGTG | GTGGATGCCG | ACGAGTACCT | 3050 |
| | CATCCCACAG | CAGGGCTTCT | TCAGCAGCCC | CTCCACGTCA | CGGACTCCCC | 3100 |
| | TCCTGAGCTC | ${\tt TCTGAGTGCA}$ | ACCAGCAACA | ATTCCACCGT | GGCTTGCATT | 3150 |
| | GATAGAAATG | GGCTGCAAAG | CTGTCCCATC | AAGGAAGACA | GCTTCTTGCA | 3200 |
| | GCGATACAGC | TCAGACCCCA | ${\tt CAGGCGCCTT}$ | GACTGAGGAC | AGCATAGACG | 3250 |
| 10 | ACACCTTCCT | ${\tt CCCAGTGCCT}$ | GAATACATAA | ACCAGTCCGT | TCCCAAAAGG | 3300 |
| | CCCGCTGGCT | CTGTGCAGAA | TCCTGTCTAT | CACAATCAGC | CTCTGAACCC | 3350 |
| | CGCGCCCAGC | AGAGACCCAC | ACTACCAGGA | CCCCACAGC | ACTGCAGTGG | 3400 |
| | GCAACCCCGA | GTATCTCAAC | ACTGTCCAGC | CCACCTGTGT | CAACAGCACA | 3450 |
| | TTCGACAGCC | ${\tt CTGCCCACTG}$ | GGCCCAGAAA | GGCAGCCACC | AAATTAGCCT | 3500 |
| 15 | GGACAACCCT | GACTACCAGC | AGGACTTCTT | TCCCAAGGAA | GCCAAGCCAA | 3550 |
| | ATGGCATCTT | TAAGGGCTCC | ACAGCTGAAA | ATGCAGAATA | CCTAAGGGTC | 3600 |
| | GCGCCACAAA | GCAGTGAATT | TATTGGAGCA | TGA | | 3630 |

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FIGURE 3C

First PCR reaction

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| EG | FR1+ EGFR3 primer | | |
|-----|--|---|-------------|
| a.) | | | ■ EGFR CDNA |
| | PCR product a encoding aa 1 to 24 fused to aa 313 to 319 | | |
| | EGFR2 + EGRR22 | 220R primer | |
| h.) | | | ■ EGFR cDNA |
| | - | PCR product b encoding aa 18 to 24 fused to aa 313 to 678 | |
| Se | cond PCR reaction | - | |
| | PCR pro | duct a + b | |
| a.) | • | extension | |
| | encoding aa | t PCR product C 1 to 24 fused to a a 313 to 678 p codon in position 679 | |
| | EGFR1 + EGFR2220R prime | er | |
| b.) | | cation of oduct c | |

FIGURE 4

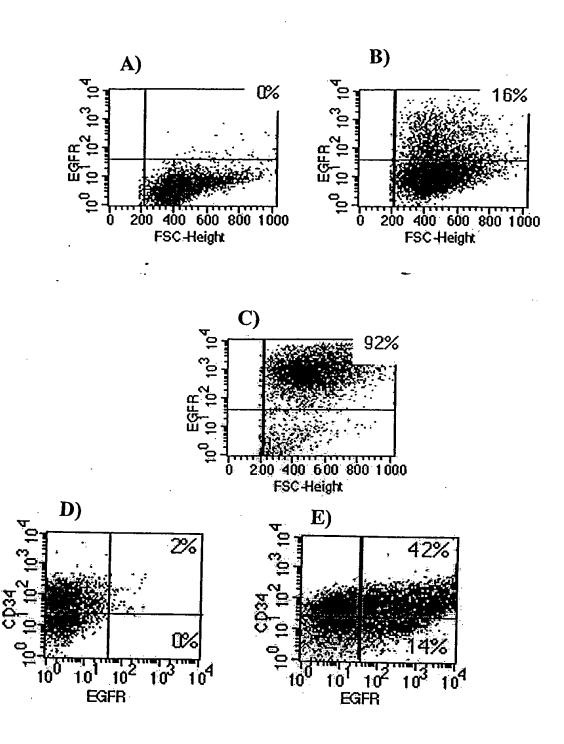


FIGURE 5

| | MRELVNIPLV | HILTLVAFSG | TEKLPKAPVI | TTPLETVDAL | VEEVATFMCA | 50 |
|----|------------|------------|-------------------|------------|------------|-----|
| | VESYPQPEIS | WTRNKILIKL | FDTRYSIREN | GQLLTILSVE | DSDDGIYCCT | 100 |
| | ANNGVGGAVE | SCGALQVKMK | PKITRPPINV | KIIEGLKAVL | PCTTMGNPKP | 150 |
| 5 | SVSWIKGDSP | LRENSRIAVL | ESGSLRIHNV | QKEDAGQYRC | VAKNSLGTAY | 200 |
| | SKVVKLEFEV | FARILRAPES | HNVTFGSFVT | LHCTATGIPV | PTITWIENGN | 250 |
| | AVSSGSIQES | VKDRVIDSRL | QLFITKPGLY | TCIATNKHGE | KFSTAKAAAT | 300 |
| | ISIAEWSKPQ | KDNKGYCAQY | RGEVCNAVLA | KDALVFLNTS | YADPEEAQEL | 350 |
| | LVHTAWNELK | VVSPVCRPAA | EALLCNHIFQ | ECSPGVVPTP | IPICREYCLA | 400 |
| 10 | VKELFCAKEW | LVMEEKTHRG | LYRSEMHLLS | VPKCSKLPSM | HWDPTACARL | 450 |
| | PHLDYNKENL | KTFPPMTSSK | PSVDIPNLPS | SSSSSFSVSP | TYSMTVIISI | 500 |
| | MSSFAIFVLL | TITTLYCCRR | RKQWKNKKRE | SAAVTLTTLP | SELLLDRLHP | 550 |
| | NPMYQRMPLL | LNPKLLSLEY | PRNNIEYVRD | IGEGAFGRVF | QARAPGLLPY | 600 |
| | EPFTMVAVKM | LKEEASADMQ | ADFQREAALM | AEFDNPNIVK | LLGVCAVGKP | 650 |
| 15 | MCLLFEYMAY | GDLNEFLRSM | SPHTVCSLSH | SDLSMRAQVS | SPGPPPLSCA | 700 |
| | EQLCIARQVA | AGMAYLSERK | FVHRDLATRN | CLVGENMVVK | IADFGLSRNI | 750 |
| | YSADYYKANE | NDAIPIRWMP | PESIFYNRYT | TESDVWAYGV | VLWEIFSYGL | 800 |
| | QPYYGMAHEE | VIYYVRDGNI | LSCPENCPVE | LYNLMRLCWS | KLPADRPSFT | 850 |
| | SIHRILERMC | ERAEGTVSV | | | | 869 |
| 20 | | | | | | |
| | | | | | | |